DOCKET NO.: ISIS-2508

PATENT

November 24, 1993 (now U.S. Patent No. 5,700,922), which is a continuation-in-part of application serial number PCT/US92/11339, filed December 23, 1992 (now U.S. Patent No. 5,623,065, issued April 22, 1997), which, in turn, is a continuation-in-part of my prior application serial number 07/814,961, filed December 24, 1991 (now abandoned). Both of the foregoing patent applications are commonly assigned with this application. –

At page 17, line 12, please insert the following paragraph before the paragraph "each of C¹-C¹ ... or heterocyclic system;"

B

--each of R^3 and R^4 is independently selected from the group consisting of hydrogen, (C_1 - C_4)alkyl, hydroxy- or alkoxy- or alkylthio-substituted (C_1 - C_4)alkyl, hydroxy, alkoxy, alkylthio and amino;--

In the Claims:

Please cancel claims 1-12, 17, 18, and 21-23. Also, please add claims 24-26, and rewrite claims 13, 19, and 20 as indicated below:

13. (amended) A method of treating an organism having a disease characterized by the undesired production of a protein, comprising contacting the organism with a macromolecule that has structure PNA-DNA-PNA and that includes a sequence of nucleobases capable of specifically hybridizing to a strand of nucleic acid coding for said protein, wherein: